

L2 ANSWER 49 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on  
STN DUPLICATE 19

ACCESSION NUMBER: 1997:483874 BIOSIS

DOCUMENT NUMBER: PREV199799783077

TITLE: RGDN peptide interaction with endothelial alpha-5-beta-1  
integrin causes sustained endothelin  
-dependent vasoconstriction of rat skeletal muscle  
arterioles.

AUTHOR(S): Mogford, Jon E.; Davis, George E.; Meininger, Gerald A.  
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CORPORATE SOURCE: Dep. Med. Physiol., Reynold's Med. Build., Texas A and M  
Univ. Health Sci. Cent., College Station, TX 77843, USA

SOURCE: Journal of Clinical Investigation, (1997) Vol. 100, No. 6,  
pp. 1647-1653.

CODEN: JCINAO. ISSN: 0021-9738.

DOCUMENT TYPE: Article

LANGUAGE: English

ENTRY DATE: Entered STN: 7 Nov 1997

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AB The ability of an integrin-binding Arg-Gly-Asp-Asn (RGDN)-containing  
peptide to influence vascular tone by interacting with the alpha-5-beta-1  
integrin was studied using rat skeletal muscle arterioles. After blockade  
of beta-3 integrin function, isolated arterioles with spontaneous tone  
showed concentration-dependent vasoconstrictions to topical application of  
GRGDNP, a peptide that shows a greater ability to interact with  
alpha-5-beta-1 than with alpha-v-beta-3. The constriction to GRGDNP (2.1  
mM) was inhibited by blocking alpha-5 integrin function, and was  
intensified by blocking beta-3 integrin function. In contrast, GRGDSP, a  
peptide that interacts better with alpha-v-beta-3, was unable to induce  
sustained constrictions. Removal of the endothelium abolished the  
vasoconstriction in response to GRGDNP, suggesting that the response was  
due to release of an endothelium-dependent factor. Indeed, blockade of  
ET-A endothelin receptors with BQ-610 (1  $\mu$ M), similar to removal of the  
endothelium and alpha-5 integrin blockade, inhibited the vasoconstriction.  
These data indicate that interaction of RGD peptides, and in particular  
the RGDN sequence with endothelial cell alpha-5-beta-1, causes  
endothelin-mediated arteriolar vasoconstriction. These results indicate  
that integrins are novel signaling receptors within the vascular wall that  
affect vasomotor tone, and may play an important role in vascular control.

TI RGDN peptide interaction with endothelial alpha-5-beta-1 integrin  
causes sustained endothelin-dependent vasoconstriction of rat  
skeletal muscle arterioles.

ACCESSION NUMBER: 96:270943 PROMT  
TITLE: Texas Biotechnology obtains endothelin A receptor  
and integrin alpha-4/beta-1 inhibitor patents  
SOURCE: BIOTECH Patent News, (1 May 1996) pp. N/A.  
ISSN: 0898-2813.  
LANGUAGE: English  
WORD COUNT: 290

\*FULL TEXT IS AVAILABLE IN THE ALL FORMAT\*

- AB Texas Biotechnology Corporation (Houston, TX; 713-796-8822) announced it was issued United States patents protecting two of its key scientific programs. Patents were issued relating to the Texas Biotechnology endothelin A receptor antagonist program as well as its VCAM/VLA-4 program.
- United States Patent 5,514,691 relates to the series of endothelin A receptor antagonists which have recently moved into pre-clinical development. This patent represents additional coverage for the Texas Biotechnology endothelin receptor antagonist program. United States Patent number 5,464,853, issued on November 7, 1995 covers composition and medicinal use for the series of Texas Biotechnology endothelin B receptor antagonists.
- The company also announced the issuance of a patent for a series of VCAM/VLA-4 inhibitors. United States Patent 5,510,332 is entitled "A process to inhibit binding of the integrin alpha-4 beta-1 VLA-4 to VCAM-1 or fibrinectin and linear peptides therefore" and was issued April 23, 1996. The integrin alpha-4/beta-1 is also referred to as VLA-4. TBC 772, a VCAM/VLA-4 inhibitor, is in late-stage research.
- Texas Biotechnology is developing acute care cardiovascular therapeutics targeting such diseases as thrombosis, congestive heart failure, chronic obstructive pulmonary disease and reperfusion injury. The lead compound, NOVASTAN (argatroban), is in Phase III clinical trials for heparin-induced thrombocytopenia at more than 100 centers in the United States. NOVASTAN is also in Phase II trials as an adjunct to thrombolytics in acute myocardial infarction at more than 75 centers in five countries. Texas Biotechnology intends to file Investigational New Drug applications for TBC 11251, an endothelin A receptor antagonist, and TBC 1269, a selectin antagonist, both small-molecule therapeutics, in 1996. The company is also generating important leads in research directed toward inhibiting cell adhesion, growth factors and programmed cell death.
- THIS IS THE FULL TEXT: COPYRIGHT 1996 BIOTECH Patent News
- TI Texas Biotechnology obtains endothelin A receptor and integrin alpha-4/beta-1 inhibitor patents

ACCESSION NUMBER: 96:178015 NLDB  
TITLE: Texas Biotechnology obtains endothelin A receptor  
and integrin alpha-4/beta-1 inhibitor patents  
SOURCE: BIOTECH Patent News, (1 May 1996) Vol. 10, No. 5.  
ISSN: 0898-2813.  
PUBLISHER: BIOTECH Patent News  
DOCUMENT TYPE: Newsletter  
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TI Texas Biotechnology obtains endothelin A receptor and  
integrin alpha-4/beta-1 inhibitor patents

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ACCESSION NUMBER: 1993:16312 BIOSIS

DOCUMENT NUMBER: PREV199344004512

TITLE: Role of integrins on endothelin  
(ET-1)-induced polymorphonuclear leukocytes (PMN) adhesion  
to the endothelium.

AUTHOR(S): Farre, A. Lopez; Riesco, A.; Espinosa, G.; Digiuni, E.;  
Alvarez, V.; Monton, M.; Gallego, M.; Casado, S.; Madrid,  
F. Sanchez; Caramelo, C.

CORPORATE SOURCE: Fundacion Jimenez Diaz, Hosp. de la Princesa, Madrid, Spain

SOURCE: Journal of the American Society of Nephrology, (1992) Vol.  
3, No. 3, pp. 629.

Meeting Info.: 25th Annual Meeting of the American Society  
of Nephrology, Baltimore, Maryland, USA, November 15-18,  
1992. J AM SOC NEPHROL.

CODEN: JASNEU. ISSN: 1046-6673.

DOCUMENT TYPE: Conference; (Meeting)

LANGUAGE: English

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TI Role of integrins on endothelin (ET-1)-induced  
polymorphonuclear leukocytes (PMN) adhesion to the endothelium.